

Andrew Freistein 10/524,993

Connecting via Winsock to STN

Welcome to STN International! Enter x:x

LOGINID:ssptabf1626

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

\* \* \* \* \* Welcome to STN International \* \* \* \* \*

NEWS 1 Web Page URLs for STN Seminar Schedule - N. America  
NEWS 2 "Ask CAS" for self-help around the clock  
NEWS 3 FEB 27 New STN AnaVist pricing effective March 1, 2006  
NEWS 4 APR 04 STN AnaVist \$500 visualization usage credit offered  
NEWS 5 MAY 10 CA/CAPLUS enhanced with 1900-1906 U.S. patent records  
NEWS 6 MAY 11 KOREAPAT updates resume  
NEWS 7 MAY 19 Derwent World Patents Index to be reloaded and enhanced  
NEWS 8 MAY 30 IPC 8 Rolled-up Core codes added to CA/CAPLUS and  
USPATFULL/USPAT2  
NEWS 9 MAY 30 The F-Term thesaurus is now available in CA/CAPLUS  
NEWS 10 JUN 02 The first reclassification of IPC codes now complete in  
INPADOC  
NEWS 11 JUN 26 TULSA/TULSA2 reloaded and enhanced with new search and  
and display fields  
NEWS 12 JUN 28 Price changes in full-text patent databases EPFULL and PCTFULL  
NEWS 13 JUL 11 CHEMSAFE reloaded and enhanced  
NEWS 14 JUL 14 FSTA enhanced with Japanese patents  
NEWS 15 JUL 19 Coverage of Research Disclosure reinstated in DWPI  
NEWS 16 AUG 09 INSPEC enhanced with 1898-1968 archive  
  
NEWS EXPRESS JUNE 30 CURRENT WINDOWS VERSION IS V8.01b, CURRENT  
MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),  
AND CURRENT DISCOVER FILE IS DATED 26 JUNE 2006.  
  
NEWS HOURS STN Operating Hours Plus Help Desk Availability  
NEWS LOGIN Welcome Banner and News Items  
NEWS IPC8 For general information regarding STN implementation of IPC 8  
NEWS X25 X.25 communication option no longer available

Enter NEWS followed by the item number or name to see news on that  
specific topic.

All use of STN is subject to the provisions of the STN Customer  
agreement. Please note that this agreement limits use to scientific  
research. Use for software development or design or implementation  
of commercial gateways or other similar uses is prohibited and may  
result in loss of user privileges and other penalties.

\* \* \* \* \* STN Columbus \* \* \* \* \*

FILE 'HOME' ENTERED AT 11:19:14 ON 16 AUG 2006

=> file reg

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

08/16/2006

Page 1

Andrew Freistein 10/524,993

	ENTRY	SESSION
FULL ESTIMATED COST	0.21	0.21

FILE 'REGISTRY' ENTERED AT 11:19:26 ON 16 AUG 2006  
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.  
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.  
COPYRIGHT (C) 2006 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file  
provided by InfoChem.

STRUCTURE FILE UPDATES: 15 AUG 2006 HIGHEST RN 901654-60-2  
DICTIONARY FILE UPDATES: 15 AUG 2006 HIGHEST RN 901654-60-2

New CAS Information Use Policies, enter HELP USAGETERMS for details.

TSCA INFORMATION NOW CURRENT THROUGH January 6, 2006

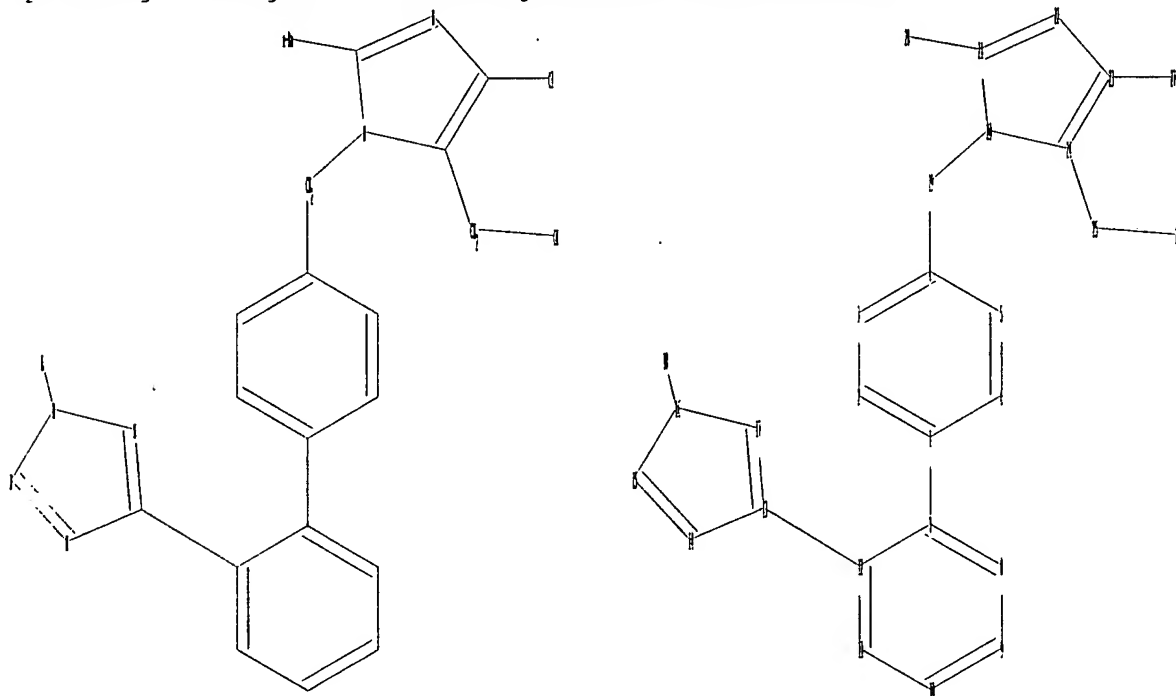
Please note that search-term pricing does apply when  
conducting SmartSELECT searches.

REGISTRY includes numerically searchable data for experimental and  
predicted properties as well as tags indicating availability of  
experimental property data in the original document. For information  
on property searching in REGISTRY, refer to:

<http://www.cas.org/ONLINE/UG/regprops.html>

=>

Uploading C:\Program Files\Stnexp\Queries\10524993\a.str



chain nodes :

18 19 25 26 27 28

ring nodes :

1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 20 21 22 23 24

Andrew Freistein 10/524,993

chain bonds :  
1-7 4-19 12-13 16-18 19-20 21-28 23-27 24-25 25-26  
ring bonds :  
1-2 1-6 2-3 3-4 4-5 5-6 7-8 7-12 8-9 9-10 10-11 11-12 13-14 13-17  
14-15 15-16 16-17 20-21 20-24 21-22 22-23 23-24  
exact/norm bonds :  
13-14 13-17 14-15 15-16 16-17 20-21 20-24 21-22 22-23 23-24  
exact bonds :  
1-7 4-19 12-13 16-18 19-20 21-28 23-27 24-25 25-26  
normalized bonds :  
1-2 1-6 2-3 3-4 4-5 5-6 7-8 7-12 8-9 9-10 10-11 11-12

Match level :  
1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom  
11:Atom 12:Atom 13:Atom 14:Atom 15:Atom 16:Atom 17:Atom 18:CLASS 19:CLASS  
20:Atom 21:Atom 22:Atom 23:Atom 24:Atom 25:CLASS 26:CLASS 27:CLASS 28:CLASS

L1 STRUCTURE UPLOADED

=> d  
L1 HAS NO ANSWERS  
L1 STR

\* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT \*

Structure attributes must be viewed using STN Express query preparation.

=> s l1  
SAMPLE SEARCH INITIATED 11:19:46 FILE 'REGISTRY'  
SAMPLE SCREEN SEARCH COMPLETED - 34 TO ITERATE

100.0% PROCESSED 34 ITERATIONS 2 ANSWERS  
SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE \*\*COMPLETE\*\*  
BATCH \*\*COMPLETE\*\*  
PROJECTED ITERATIONS: 331 TO 1029  
PROJECTED ANSWERS: 2 TO 124

L2 2 SEA SSS SAM L1

=> s l1 full  
FULL SEARCH INITIATED 11:19:49 FILE 'REGISTRY'  
FULL SCREEN SEARCH COMPLETED - 673 TO ITERATE

100.0% PROCESSED 673 ITERATIONS 34 ANSWERS  
SEARCH TIME: 00.00.01

L3 34 SEA SSS FUL L1

=> file caplus  
COST IN U.S. DOLLARS SINCE FILE TOTAL  
ENTRY SESSION  
FULL ESTIMATED COST 166.94 167.15

FILE 'CAPLUS' ENTERED AT 11:20:05 ON 16 AUG 2006  
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.  
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.  
COPYRIGHT (C) 2006 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 16 Aug 2006 VOL 145 ISS 8  
FILE LAST UPDATED: 15 Aug 2006 (20060815/ED)

Effective October 17, 2005, revised CAS Information Use Policies apply. They are available for your review at:

<http://www.cas.org/infopolicy.html>

=> s 13

L4 2811 L3

=> s 14 and polymorph? or cyrstalline

191744 POLYMORPH?

21 CYRSTALLINE

L5 68 L4 AND POLYMORPH? OR CYRSTALLINE

=> s losartan

4698 LOSARTAN

1 LOSARTANS

L6 4698 LOSARTAN  
(LOSARTAN OR LOSARTANS)

=> s polymorph or crystal?

7423 POLYMORPH

8513 POLYMORPHS

12981 POLYMORPH

(POLYMORPH OR POLYMORPHS)

1765010 CRYSTAL?

345195 CRYST

1801 CRYSTS

346463 CRYST

(CRYST OR CRYSTS)

89394 CRYSTD

18928 CRYSTG

232964 CRYSTN

2373 CRYSTNS

234266 CRYSTN

(CRYSTN OR CRYSTNS)

2064318 CRYSTAL?

(CRYSTAL? OR CRYST OR CRYSTD OR CRYSTG OR CRYSTN)

L7 2067837 POLYMORPH OR CRYSTAL?

=> s 17 and 16

L8 57 L7 AND L6

=> s 18 and potassium

597707 POTASSIUM

16 POTASSIUMS

597709 POTASSIUM

(POTASSIUM OR POTASSIUMS)

L9 27 L8 AND POTASSIUM

=> d scan

L9 27 ANSWERS CAPLUS COPYRIGHT 2006 ACS on STN

IC ICM A61K031-4178

ICS A61P009-12; C07D403-10

CC 63-6 (Pharmaceuticals)

Section cross-reference(s): 28

TI Purification methods in preparation of pharmaceutical salts of losartan

ST pharmaceutical salt losartan property prepn

IT Crystal structure  
(of losartan salts)

IT Density

Dissolution

Flow

Freeze drying

Particle size distribution

Polymorphism (crystal)

Solubility

Stability

Surface area

(purification methods in preparation of pharmaceutical salts of losartan)

IT Alcohols, processes

RL: PEP (Physical, engineering or chemical process); PYP (Physical process); PROC (Process)

(purification methods in preparation of pharmaceutical salts of losartan)

IT Alkali metal hydroxides

Alkali metals, reactions

Alkaline earth hydroxides

Alkaline earth metals

RL: RCT (Reactant); RACT (Reactant or reagent)

(purification methods in preparation of pharmaceutical salts of losartan)

IT Drug delivery systems

(tablets, coated; purification methods in preparation of pharmaceutical salts of losartan)

IT 9004-34-6, Cellulose, biological studies

RL: PEP (Physical, engineering or chemical process); PYP (Physical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

(microcryst.; purification methods in preparation of pharmaceutical salts of losartan)

IT 60-29-7, Diethyl ether, processes 67-56-1, Methanol, processes

67-63-0, 2-Propanol, processes 141-78-6, EtOAc, processes 142-82-5, Heptane, processes

RL: PEP (Physical, engineering or chemical process); PYP (Physical process); PROC (Process)

(purification methods in preparation of pharmaceutical salts of losartan)

)  
 IT 7631-86-9, Silica, biological studies 64044-51-5 74811-65-7,  
 Croscarmellose sodium  
 RL: PEP (Physical, engineering or chemical process); PYP (Physical  
 process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);  
 USES (Uses)  
 (purification methods in preparation of pharmaceutical salts of losartan  
 )  
 IT 114798-26-4DP, Losartan, salts 124750-99-8P, Losartan  
 potassium 733047-57-9P 733047-58-0P 733047-59-1P  
 RL: PRP (Properties); PUR (Purification or recovery); SPN (Synthetic  
 preparation); THU (Therapeutic use); BIOL (Biological study); PREP  
 (Preparation); USES (Uses)  
 (purification methods in preparation of pharmaceutical salts of losartan  
 )  
 IT 865-47-4 865-48-5 1305-62-0, Calcium hydroxide, reactions 2414-98-4,  
 Magnesium ethoxide 79047-41-9 133051-88-4  
 RL: RCT (Reactant); RACT (Reactant or reagent)  
 (purification methods in preparation of pharmaceutical salts of losartan  
 )  
 IT 114798-26-4P, Losartan 133909-99-6P  
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT  
 (Reactant or reagent)  
 (purification methods in preparation of pharmaceutical salts of losartan  
 )

HOW MANY MORE ANSWERS DO YOU WISH TO SCAN? (1):0

=> d ibib abs 1-27

L9 ANSWER 1 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN  
 ACCESSION NUMBER: 2006:699729 CAPLUS  
 DOCUMENT NUMBER: 145:152705  
 TITLE: Stable noncrystalline formulations comprising  
 losartan  
 INVENTOR(S): Palakodaty, Srinivas; Kordikowski, Andreas; Daintree,  
 Linda Sharon; Duddu, Sarma; Kugler, Alan; Zhang,  
 Jiang; Snyder, Herman; Lechuga, David; Palepu, Nagesh;  
 Eldon, Michael A.  
 PATENT ASSIGNEE(S): Nektar Therapeutics, USA  
 SOURCE: PCT Int. Appl., 96 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2006076097	A2	20060720	WO 2005-US44278	20051206
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW			
RW:	AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH,			

GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,  
 KG, KZ, MD, RU, TJ, TM  
 US 2006160871 A1 20060720 US 2005-296108 20051206  
 PRIORITY APPLN. INFO.: US 2004-633988P P 20041207  
 AB One or more embodiments of the invention provide various novel  
 formulations, and tablet dosage forms, comprising losartan that  
 are noncryst., stable, and/or otherwise improvements over known  
 losartan formulations. One or more embodiments of the invention  
 further provide methods for preparing the formulation, methods for preparing  
 the  
 tablet dosage form, and to methods of administering the tablet dosage  
 and/or formulation comprising losartan. The losartan  
 -containing formulations may be administered to a user to treat hypertension,  
 and related conditions. A spray drying process is used to produce  
 particles comprising non-crystalline losartan and a  
 stabilizing excipient. The stabilizing excipient comprises a copolymer,  
 such as a vinyl pyrrolidone-vinyl acetate copolymer.

L9 ANSWER 2 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN  
 ACCESSION NUMBER: 2006:430764 CAPLUS  
 DOCUMENT NUMBER: 145:46064  
 TITLE: Process for preparation of H-type crystalline  
 form of losartan potassium  
 INVENTOR(S): Wang, Youhu; Zhou, Minghua; Hu, Gongyun; Wang, Danhua;  
 Jin, Yongjun; Chai, Jian; Li, Wei  
 PATENT ASSIGNEE(S): Zhejiang Huahai Pharmaceutical Co., Ltd., Peop. Rep.  
 China  
 SOURCE: Faming Zhuanli Shenqing Gongkai Shuomingshu, 15 pp.  
 CODEN: CNXXEV  
 DOCUMENT TYPE: Patent  
 LANGUAGE: Chinese  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
CN 1763036	A	20060426	CN 2004-10067407	20041022
PRIORITY APPLN. INFO.:			CN 2004-10067407	20041022
AB The title crystalline form of losartan potassium features in having characteristic absorption peak 20 of 7.11, 7.32, 11.08, 14.24, 14.80, 18.55, 18.93, 21.38, 23.90, 28.72, 29.88, 30.97, 33.11 according to XRD pattern, and exhibiting maximum endothermic fusion peak at 273.37°C, maximum endothermic onset temperature of crystalline transition of 235.47°C under detection condition of 30-300°C, heating rate of 10°C/min, open, and nitrogen gas flow of 40 mL/min according to DSC thermogram. The title method comprises dissolving losartan potassium in alc. optionally containing water 1-15%, adding mixture of hydrocarbon/alc. with volume ratio of 10:1-20:1, cooling to 0-30°C, stirring for 0.5-1 h, standing for 1-3 h, separating, and drying at 20-120°C to obtain the final product.				

L9 ANSWER 3 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN  
 ACCESSION NUMBER: 2006:343149 CAPLUS  
 DOCUMENT NUMBER: 144:370099  
 TITLE: Alkylation and reduction process for preparation of  
 2-butyl-4-chloro-1-[[2'-(2-triphenylmethyl-2H-tetrazol-  
 5-yl)-1,1'-biphenyl-4-yl]methyl]-1H-imidazole-5-  
 methanol which is a losartan intermediate  
 INVENTOR(S): Chava, Satyanaryana; Vasireddy, Umamaheswar Rao;  
 Vellanki, Siva Ram Prasad; Balusu, Rajababu

PATENT ASSIGNEE(S): Matrix Laboratories Ltd., India  
 SOURCE: PCT Int. Appl., 12 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
-----	----	-----	-----	-----
WO 2006038223	A1	20060413	WO 2005-IN308	20050913
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				

PRIORITY APPLN. INFO.: IN 2004-CH1032 A 20041006

OTHER SOURCE(S): CASREACT 144:370099

AB 2-Butyl-4-chloro-1-[[2'-(2-triphenylmethyl-2H-tetrazol-5-yl)-1,1'-biphenyl-4-yl]methyl]-1H-imidazole-5-methanol (I), an intermediate in the synthesis of losartan and its pharmaceutically acceptable salts, is prepared in high yield and selectivity by: (A) the alkylation of 2-butyl-4-chloroimidazole-5-carboxaldehyde with 5-(4'-bromomethyl-1,1'-biphenyl-2-yl)-1-(triphenylmethyl)-1H-tetrazole in water and a haloalkane (e.g., 1-chlorobutane) in the presence of a base (e.g., NaOH) and a phase-transfer catalyst (e.g., tetrabutylammonium bromide); (B) maintaining the reaction mass at 10-65° for 18-48 h; (C) letting the reaction mixture settle and separating the aqueous and organic layers; (D) reacting the organic layer with sodium borohydride and a lower alc. (e.g., methanol); (E) adding water to the reaction mass and separating the layers; (F) washing the organic layer with water; (G) concentrating the organic layer; (H) isolating the I and drying it at 45-75°; and (I) conducting an optional crystallization of I if necessary.

REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 4 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2005:1004564 CAPLUS  
 DOCUMENT NUMBER: 143:292576  
 TITLE: Stabilization of a polymorphic form of losartan potassium  
 INVENTOR(S): Svete, Peter; Grahek, Rok; Humar, Vlasta; Husu-Kovacevic, Breda; Jerala-Strukelj, Zdenka  
 PATENT ASSIGNEE(S): Lek Pharmaceuticals D.D., Slovenia  
 SOURCE: PCT Int. Appl., 22 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 2  
 PATENT INFORMATION:



PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2005084670	A1	20050915	WO 2005-EP2108	20050228
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				

PRIORITY APPLN. INFO.: SI 2004-67 A 20040301

AB Comps. were developed which stabilize an active pharmaceutical ingredient in polymorph form susceptible to degradation or interconversion into other polymorph forms, where stabilizing substance is conveniently among silicon dioxide, silicified microcryst. cellulose, magnesium oxide and polyethylene glycol. The polymorphic form of losartan potassium was stable when formulated with Syloid and PEG 6000.

REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 5 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2005:967720 CAPLUS

DOCUMENT NUMBER: 144:181122

TITLE: Losartan potassium 3.5-hydrate, a new crystalline form

AUTHOR(S): Hu, Xiu Rong; Wang, Yun Wu; Gu, Jian Ming

CORPORATE SOURCE: Center of Analysis and Measurement, Zhejiang University, Zhejiang, 310028, Peop. Rep. China

SOURCE: Acta Crystallographica, Section E: Structure Reports Online (2005), E61(9), m1686-m1688  
 CODEN: ACSEBH; ISSN: 1600-5368  
 URL: <http://journals.iucr.org/e/issues/2005/09/00/dn6239/dn6239Isup2.hkl>

PUBLISHER: Blackwell Publishing Ltd.

DOCUMENT TYPE: Journal; (online computer file)

LANGUAGE: English

AB Crystals of the title compound are orthorhombic, space group Pbca, with a 13.1389(3), b 25.6885(5), c 31.1822(7) Å; Z = 8, dc = 1.323; R = 0.054, Rw(F2) = 0.124 for 6573 reflections. The asym. unit is composed of two losartan anions, two K<sup>+</sup> cations and seven H<sub>2</sub>O mols. Some H<sub>2</sub>O mols. bridge the K ions linking the mols. to form an infinite chain. The two K ions have different environments; one is six-coordinated by three water O atoms and three tetrazole N atoms, whereas the other is five-coordinated by five water O atoms. Extensive H-bonding interactions lead to a three-dimensional structure.

REFERENCE COUNT: 13 THERE ARE 13 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 6 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2005:638868 CAPLUS

DOCUMENT NUMBER: 143:139094

TITLE: An improved process for the synthesis of losartan potassium

INVENTOR(S): Kumar, Ashok; Singh, Rajesh Kumar Keshava Prasad; Panda, Nalinakshya Balaram; Upare, Abhay Atmaram;

Nimbalkar, Manmohan Madhavrao; Soudagar, Satish  
 Rajanikant; Saxena, Ashvini Kumar Nand Kishore  
 PATENT ASSIGNEE(S): Ipca Laboratories Limited, India  
 SOURCE: PCT Int. Appl., 23 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2005066158	A2	20050721	WO 2004-IN169	20040615
WO 2005066158	A3	20050825		

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW  
 RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

PRIORITY APPLN. INFO.: IN 2004-MU14 A 20040106  
 AB Improved processes using primary, secondary and tertiary alcs. and with safer mode of introduction of the reagent and reaction conditions are described. Further, the process of manufacture of losartan potassium by use of alkali metal salt such as potassium carbonate is disclosed. A process for preparation of the polymorphic Form I of losartan potassium is also disclosed herein. Losartan potassium was prepared by the reaction of potassium tertiary butoxide with trityl losartan in methanol, yield = 78%.

L9 ANSWER 7 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN  
 ACCESSION NUMBER: 2005:527406 CAPLUS  
 DOCUMENT NUMBER: 143:65424  
 TITLE: Process for the preparation of crystalline losartan potassium  
 INVENTOR(S): Razzetti, Gabriele; Magrone, Domenico; Ercoli, Mauro; Allegrini, Pietro; Castaldi, Graziano  
 PATENT ASSIGNEE(S): Dipharma S.P.A., Italy  
 SOURCE: U.S. Pat. Appl. Publ., 4 pp.  
 CODEN: USXXCO  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2005131040	A1	20050616	US 2004-10455	20041214
EP 1544198	A1	20050622	EP 2004-27598	20041119

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK, HR, IS, YU  
 JP 2005179355 A2 20050707 JP 2004-360875 20041214  
 PRIORITY APPLN. INFO.: IT 2003-MI2472 A 20031216

AB A process for the preparation of crystalline losartan potassium and crystalline hydrate losartan potassium is disclosed. A suspension of losartan in EtOAc was treated with KHC03 in water to give losartan potassium crystalline hydrate.

L9 ANSWER 8 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2005:284148 CAPLUS

DOCUMENT NUMBER: 142:341839

TITLE: Process for the synthesis of losartan potassium

INVENTOR(S): Kumar, Ashok; Singh, Rajesh Kumar Keshava Prasad; Panda, Nalinakshya Balaram; Upare, Abhay Atmaram; Nimbalkar, Manmohan Madhavrao; Soudagar, Satish Rajanikant; Saxena, Ashvini Kumar Nand Kishore

PATENT ASSIGNEE(S): India

SOURCE: U.S. Pat. Appl. Publ., 12 pp., Cont.-in-part of U.S. Ser. No. 431,847.

CODEN: USXXCO

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2005070586	A1	20050331	US 2004-913121	20040805
US 2004224998	A1	20041111	US 2003-431847	20030507
US 6916935	B2	20050712		

PRIORITY APPLN. INFO.: US 2003-431847 A2 20030507  
US 2003-468208P P 20030506

AB Improved processes using primary, secondary and tertiary alcs. and with safer mode of introduction of the reagent and reaction conditions are described. Further, the process of manufacture of losartan potassium by use of alkali metal salt such as potassium carbonate is disclosed. A process for preparation of the polymorphic form I of losartan potassium is also disclosed herein. Potassium tertiary butoxide was reacted with trityl losartan in methanol to obtain losartan potassium form I, yield: 78%.

L9 ANSWER 9 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2005:280278 CAPLUS

DOCUMENT NUMBER: 143:163500

TITLE: 4'-(2-Butyl-4-chloro-5-formylimidazol-1-ylmethyl)biphenyl-2-carbonitrile

AUTHOR(S): Yathirajan, Hemmige S.; Nagaraj, Basavegowda; Narasegowda, Rajenahally S.; Nagaraja, Padmarajaiah; Bolte, Michael

CORPORATE SOURCE: Department of Studies in Chemistry, University of Mysore, Manasagangotri, Mysore, 570 006, India

SOURCE: Acta Crystallographica, Section E: Structure Reports Online (2005), E61(4), o1193-o1195  
CODEN: ACSEBH; ISSN: 1600-5368

URL: <http://journals.iucr.org/e/issues/2005/04/00/dn6209/index.html>

PUBLISHER: Blackwell Publishing Ltd.

DOCUMENT TYPE: Journal; (online computer file)

LANGUAGE: English

AB The title compound, C22H20ClN3O, (I), was used as an intermediate for the

synthesis of the antihypertensive drug losartan.  
Crystallog. data are given. Bond lengths and angles are  
unexceptional. The crystal packing is stabilized by one  
C-H...O and one C-H...N contact.

It is noteworthy that (I) is isomorphous with a closely related compound  
which differs in having a but-2-enyl chain instead of a Bu chain on the  
imidazole ring.

REFERENCE COUNT: 10 THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS  
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 10 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:964836 CAPLUS

DOCUMENT NUMBER: 141:400944

TITLE: Losartan potassium synthesis

INVENTOR(S): Kumar, Ashok; Singh, Rajeshkumar; Panda, Nalinakshya;  
Upare, Abhay; Nimbalkar, Manmohan; Soudagar, Satish

PATENT ASSIGNEE(S): Ipca Laboratories, India

SOURCE: U.S. Pat. Appl. Publ., 5 pp.

CODEN: USXXCO

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2004224998	A1	20041111	US 2003-431847	20030507
US 6916935	B2	20050712		
US 2005070586	A1	20050331	US 2004-913121	20040805
US 2005043539	A1	20050224	US 2004-938317	20040910
PRIORITY APPLN. INFO.:			US 2003-468208P	P 20030506
			US 2003-431847	A2 20030507
			IN 2004-MU80	A 20040128

AB A process for the preparation of losartan potassium (I) by  
reacting trityllosartan in a primary alc. with a potassium  
tertiary alkoxide is disclosed. I was prepared in 81.63% yield by refluxing  
a solution of potassium tert-butoxide in MeOH with trityllosartan  
for 8 h.

REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS  
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 11 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:930812 CAPLUS

DOCUMENT NUMBER: 142:288051

TITLE: Trityl losartan

AUTHOR(S): Sieron, Leslaw; Nagaraj, B.; Prabhuswamy, B.;  
Yathirajan, H. S.; Nagaraja, P.; Narasegowda, R. S.;  
Gaonkar, S. L.

CORPORATE SOURCE: Institute of General and Ecological Chemistry,  
Technical University of Lodz, Lodz, 90-924, Pol.

SOURCE: Acta Crystallographica, Section C: Crystal Structure  
Communications (2004), C60(11), o821-o823  
CODEN: ACSCEE; ISSN: 0108-2701

PUBLISHER: Blackwell Publishing Ltd.

DOCUMENT TYPE: Journal

LANGUAGE: English

AB The title compound (systematic name: {2-butyl-4-chloro-1-[2'-(2-trityl-2H-  
tetrazol-5-yl)biphenyl-4-ylmethyl]-1H-imidazol-5-yl}methanol),  
C41H37ClN6O, crystallizes in the centrosym. space group  
P.hivin.1 with two independent mols. in the asym. unit.

Crystallog. data are given. These mols. differ significantly only in the relative orientations of the rings in the biphenyltetrazole moieties. One of the mols. shows disorder for three C atoms in the Bu group. H bonds link the mols. in an infinite chain along the a axis.

REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 12 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:857589 CAPLUS

DOCUMENT NUMBER: 141:337645

TITLE: A process for the synthesis of losartan potassium

INVENTOR(S): Kumar, Ashok; Singh, Rajesh Kumar; Panda, Nalinakshya; Upare, Abhay Atmaram; Nimbalkar, Manmohan Madhavrao; Soudagar, Satish Rajanikant

PATENT ASSIGNEE(S): Ipca Laboratories Limited, India

SOURCE: PCT Int. Appl., 13 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004087691	A1	20041014	WO 2003-IN230	20030627
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW			
RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG			
AU 2003269470	A1	20041025	AU 2003-269470	20030627
EP 1608641	A1	20051228	EP 2003-751250	20030627
R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK			
PRIORITY APPLN. INFO.:			IN 2003-MU335	A 20030403
			WO 2003-IN230	W 20030627

AB A process for the synthesis of losartan potassium comprises reacting trityl losartan in a primary alc. with potassium tertiary alkoxide. A solution of potassium tertiary butoxide in methanol was refluxed with trityl losartan under N for 8 h to obtain losartan potassium which was separated and purified (yield=81.63%).

REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 13 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:740318 CAPLUS

DOCUMENT NUMBER: 141:248755

TITLE: Amorphous form of losartan potassium

INVENTOR(S): Parthasaradhi, Reddy Bandi; Rathnakar, Reddy Kura; Raji, Reddy Rapolu; Narasa, Reddy Attunuri; Narasa, Reddy Bolla

PATENT ASSIGNEE(S): Hetero Drugs Limited, India

SOURCE: PCT Int. Appl., 8 pp.

CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004076443	A1	20040910	WO 2003-IN37	20030225
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
AU 2003209669	A1	20040917	AU 2003-209669	20030225
PRIORITY APPLN. INFO.: WO 2003-IN37 A 20030225 AB The invention relates to a novel amorphous form of losartan potassium, to a process for the preparation thereof, and to a composition containing it. Losartan potassium crystals (50 g) were added to a mixture containing MeOH and EtOAc and the solution was dried for 18 h to give 42 g losartan potassium. REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT				

L9 ANSWER 14 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN  
 ACCESSION NUMBER: 2004:740317 CAPLUS  
 DOCUMENT NUMBER: 141:265973  
 TITLE: Preparation of polymorphic crystal forms of  
 the antihypertensive agent losartan  
 potassium  
 INVENTOR(S): Kumar, Pananchukunnath Manoj; Manikandan, Ramalingam;  
 Singh, Romi Barat; Nagaprasad, Vishnubhotla; Malik,  
 Rajiv  
 PATENT ASSIGNEE(S): Ranbaxy Laboratories Limited, India  
 SOURCE: PCT Int. Appl., 22 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004076442	A1	20040910	WO 2004-IB516	20040227
W: AE, AE, AG, AL, AL, AM, AM, AM, AT, AT, AU, AZ, AZ, BA, BB, BG, BG, BR, BR, BW, BY, BY, BZ, BZ, CA, CH, CN, CN, CO, CO, CR, CR, CU, CU, CZ, CZ, DE, DE, DK, DK, DM, DZ, EC, EC, EE, EE, EG, ES, ES, FI, FI, GB, GD, GE, GE, GH, GM, HR, HR, HU, HU, ID, IL, IN, IS, JP, JP, KE, KE, KG, KG, KP, KP, KP, KR, KR, KZ, KZ, KZ, LC, LK, LR, LS, LS, LT, LU, LV, MA, MD, MD, MG, MK, MN, MW, MX, MX, MZ, MZ, NA, NI RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN,				

GQ, GW, ML, MR, NE, SN, TD, TG, BF, BJ, CF, CG, CI, CM, GA, GN,  
GQ, GW, ML, MR, NE, SN, TD, TG

PRIORITY APPLN. INFO.: IN 2003-DE202 A 20030228

AB Processes for producing polymorphic crystal forms of losartan potassium (I), useful as an antihypertensive, are claimed as are the crystal polymorphs of I, their crystal-characterization data, and their use in pharmaceutical formulations.

REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 15 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:740288 CAPLUS

DOCUMENT NUMBER: 141:248753

TITLE: Preparation of losartan potassium polymorphs

INVENTOR(S): Boccignone, Andrea; Malpezzi, Luciana; Castaldi, Graziano; Allegrini, Pietro; Beltrame, Andrea

PATENT ASSIGNEE(S): Dinamite Dipharma S.P.A. In Abbreviate Form Dipharma S.P.A., Italy; Dipharma S.P.A.

SOURCE: PCT Int. Appl., 25 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004076406	A2	20040910	WO 2004-EP1717	20040220
WO 2004076406	A3	20050113		
W:	AE, AE, AG, AL, AL, AM, AM, AM, AT, AT, AU, AZ, AZ, BA, BB, BG, BG, BR, BR, BW, BY, BY, BZ, BZ, CA, CH, CN, CN, CO, CO, CR, CR, CU, CU, CZ, CZ, DE, DE, DK, DK, DM, DZ, EC, EC, EE, EE, EG, ES, ES, FI, FI, GB, GD, GE, GE, GH, GM, HR, HR, HU, HU, ID, IL, IN, IS, JP, JP, KE, KE, KG, KG, KP, KP, KR, KR, KZ, KZ, KZ, LC, LK, LR, LS, LS, LT, LU, LV, MA, MD, MD, MG, MK, MN, MW, MX, MX, MZ, MZ, NA, NI			
RW:	BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG			

PRIORITY APPLN. INFO.: IT 2003-MI328 A 20030225

AB Losartan potassium polymorphs, identified as losartan potassium crystalline hydrate, losartan potassium amorphous and losartan potassium modification crystalline III, a process for their preparation, pharmaceutical compns. containing them and their use in therapy. Thus, losartan was dissolved in MeOH and treated with KHCO<sub>3</sub> to give a losartan potassium polymorph III.

L9 ANSWER 16 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:648382 CAPLUS

DOCUMENT NUMBER: 141:179636

TITLE: Purification methods in preparation of pharmaceutical salts of losartan

INVENTOR(S): Antoncic, Ljubomir; Copar, Anton; Svete, Peter; Husu-Kovacecic, Breda; Ham, Zoran; Marolt, Boris

PATENT ASSIGNEE(S): Lek Pharmaceuticals D.D., Slovenia

SOURCE: PCT Int. Appl., 110 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004066997	A2	20040812	WO 2004-SI1	20040129
WO 2004066997	A3	20041111		
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI			
SI 21423	C	20040831	SI 2003-25	20030130
SI 21424	C	20040831	SI 2003-26	20030130
SI 21508	C	20041231	SI 2003-145	20030612
SI 21509	C	20041231	SI 2003-157	20030626
EP 1589966	A2	20051102	EP 2004-706411	20040129
R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK			
US 2006004207	A1	20060105	US 2005-524993	20050218
PRIORITY APPLN. INFO.:			SI 2003-25	A 20030130
			SI 2003-26	A 20030130
			SI 2003-145	A 20030612
			SI 2003-157	A 20030626
			SI 2003-270	A 20031105
			WO 2004-SI1	W 20040129

AB Pharmaceutical crystalline and amorphous alkali and alkaline earth salts of losartan are prepared and new manufacturing, purification and isolation procedures for the salts in high purity are disclosed. Stable pharmaceutical compns. containing new crystalline potassium salts of losartan were prepared. Thus, losartan potassium was prepared from losartan by treatment with KOH in iso-PrOH. Film-coated tablets contained losartan potassium 100.000 mg.

L9 ANSWER 17 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN  
 ACCESSION NUMBER: 2004:589438 CAPLUS  
 DOCUMENT NUMBER: 141:111635  
 TITLE: Body weight gain inhibitors containing angiotensin II antagonists  
 INVENTOR(S): Terashita, Zen-ichi; Kusumoto, Keiji; Yamaguchi, Fuminari; Imura, Yoshimi  
 PATENT ASSIGNEE(S): Takeda Chemical Industries, Ltd., Japan  
 SOURCE: PCT Int. Appl., 54 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: Japanese  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004060399	A1	20040722	WO 2003-JP16656	20031225
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ,			



OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM,  
 TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW  
 RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,  
 BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,  
 ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK,  
 TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

CA 2511737 AA 20040722 CA 2003-2511737 20031225  
 AU 2003292775 A1 20040729 AU 2003-292775 20031225  
 JP 2004217648 A2 20040805 JP 2003-429424 20031225  
 EP 1579872 A1 20050928 EP 2003-768195 20031225

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,  
 IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK

CN 1756567 A 20060405 CN 2003-80110043 20031225  
 US 2006069133 A1 20060330 US 2005-540369 20050623

PRIORITY APPLN. INFO.: JP 2002-380386 A 20021227  
 WO 2003-JP16656 W 20031225

OTHER SOURCE(S): MARPAT 141:111635

AB It is intended to provide a drug showing an excellent effect of inhibiting body weight gain which contains a compound having an angiotensin II antagonism, its prodrug or a salt thereof. It is also intended to provide a drug capable of inhibiting body weight gain in a patient even in the case of administering a therapeutically efficacious PPAR $\gamma$ -agonistic substance in treating diabetes or other diseases. The effect of 2-ethoxy-1-[[2'-(2,5-dihydro-5-oxo-1,2,4-oxadiazol-3-yl)biphenyl-4-yl]methyl]benzimidazol-7-carboxylic acid (I) on PPAR $\gamma$ -agonistic substance (pioglitazone)-induced obesity in rats was examined. A capsule containing I 5, pioglitazone hydrochloride 30, lactose 85, fine cryst. cellulose 70, magnesium stearate 10 mg was formulated.

L9 ANSWER 18 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:414643 CAPLUS

DOCUMENT NUMBER: 140:412339

TITLE: Crystalline form of losartan potassium

INVENTOR(S): Reddy, Manne Satyanarayana; Eswaraiah, Sajja; Koppera, Ravinder Reddy; Reddy, Vajrала Venkata

PATENT ASSIGNEE(S): Reddy's Laboratories Limited, India; Reddy's Laboratories, Inc.

SOURCE: U.S. Pat. Appl. Publ., 11 pp.

CODEN: USXXCO

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2004097568	A1	20040520	US 2003-629316	20030729
PRIORITY APPLN. INFO.:			IN 2002-MA568	A 20020729

AB A compound that is a crystalline Form III of losartan potassium is provided. Also provided are compns. containing the compound and methods for its preparation. For example, 125 g of trityl losartan (preparation given) was mixed with an aqueous solution containing 11 g of KOH, 125 mL water, and 1250 mL methanol until the reaction was complete. The solvent was distilled off the reaction solution under vacuum, and water (325 mL) added to the residual mass, stirred for 30 min, the pH adjusted to 8.2 to 8.8, and the mass filtered. The filtrate was washed with water, the water was distilled off, and the resulting residue was dissolved in methanol,

the solvent distilled off, and the residual mass cooled to a temperature of 5 to

10°, filtered, and dried to yield crystalline polymorph Form III of losartan potassium (weight 43.0 g). The crystalline polymorph Form III of losartan potassium was also obtained from crystalline polymorph Form I of losartan potassium.

L9 ANSWER 19 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2003:454301 CAPLUS

DOCUMENT NUMBER: 139:26612

TITLE: Amorphous and crystalline forms of losartan potassium

INVENTOR(S): Dolitzky, Ben Zion; Weizel, Shlomit; Nisnevich, Gennady; Rukhman, Igor; Kaftanov, Julia

PATENT ASSIGNEE(S): Teva Pharmaceutical Industries, Ltd., Israel; Teva Pharmaceuticals USA, Inc.

SOURCE: PCT Int. Appl., 46 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2003048135	A1	20030612	WO 2002-US36550	20021113
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW			
RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG			
CA 2465597	AA	20030612	CA 2002-2465597	20021113
AU 2002360386	A1	20030617	AU 2002-360386	20021113
US 2004006237	A1	20040108	US 2002-293820	20021113
EP 1458693	A1	20040922	EP 2002-795637	20021113
R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, SK			
CN 1612866	A	20050504	CN 2002-826988	20021113
JP 2006504618	T2	20060209	JP 2003-549327	20021113
ZA 2004003582	A	20050511	ZA 2004-3582	20040511
NO 2004002434	A	20040611	NO 2004-2434	20040611
PRIORITY APPLN. INFO.:			US 2001-333034P	P 20011114
			US 2002-401278P	P 20020805
			WO 2002-US36550	W 20021113

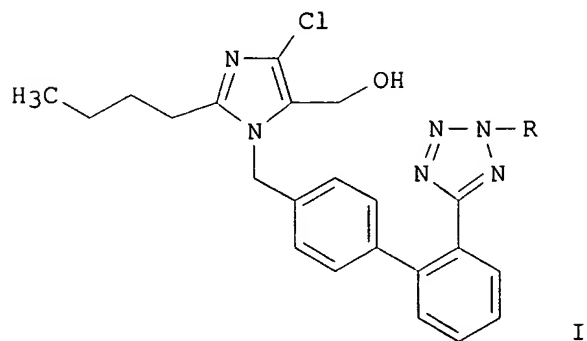
AB This invention relates to novel amorphous losartan potassium, novel losartan potassium in a crystalline form that is a hydrate, novel crystalline losartan potassium Form IV and solvates thereof, novel crystalline losartan potassium Form V and solvates thereof, to processes for their preparation, to compns. containing them and to their use in medicine. This invention further relates to a novel process for preparing crystalline losartan potassium Form I and Form II.

REFERENCE COUNT: 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS

## RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 20 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN  
 ACCESSION NUMBER: 2002:906207 CAPLUS  
 DOCUMENT NUMBER: 138:4604  
 TITLE: Deprotection process for the crystallization  
 of losartan potassium in the  
 polymorphic crystalline form I  
 INVENTOR(S): Ramashankar; Reddy, Ravinder Vennapu; Sivakumaran,  
 Meenakshisunderam; Handa, Vijay Kumar  
 PATENT ASSIGNEE(S): Aurobindo Pharma Limited, India  
 SOURCE: PCT Int. Appl., 17 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002094816	A1	20021128	WO 2001-IN205	20011120
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
IN 193625	A	20040731	IN 2001-MA403	20010518
EP 1294712	A1	20030326	EP 2001-274254	20011120
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR				
SI 21236	C	20031231	SI 2001-20042	20011120
JP 2004520446	T2	20040708	JP 2002-591489	20011120
BG 107478	A	20040130	BG 2003-107478	20030117
PRIORITY APPLN. INFO.:			IN 2001-MA403	A 20010518
			IN 2001-CH403	A 20010518
			WO 2001-IN205	W 20011120
OTHER SOURCE(S):			CASREACT 138:4604; MARPAT 138:4604	
GI				



AB The polymorphic crystalline form I of losartan

potassium (I; R = K) is prepared in high yield and selectivity by the deprotection of a losartan precursor (I; R = H, CPh<sub>3</sub>; e.g., trityl losartan) with potassium hydroxide in an alc. (e.g., methanol), followed by reducing the alc. concentration under vacuum, and adding a nonsolvent (e.g., acetone) to precipitate the losartan potassium.

REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 21 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2002:496878 CAPLUS

DOCUMENT NUMBER: 137:286738

TITLE: Losartan potassium, a non-peptide agent for the treatment of arterial hypertension

AUTHOR(S): Fernandez, Daniel; Vega, Daniel; Ellena, Javier A.; Echeverria, Gustavo

CORPORATE SOURCE: Escuela de Ciencia y Tecnologia, Universidad Nacional de General San Martin, Buenos Aires, Argent.

SOURCE: Acta Crystallographica, Section C: Crystal Structure Communications (2002), C58(7), m418-m420  
CODEN: ACSCEE; ISSN: 0108-2701

PUBLISHER: Blackwell Munksgaard

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Crystals of the title compound, potassium 2-butyl-4-chloro-1-([2'-(5-tetrazolido)biphenyl-4-yl]methyl)-1H-imidazol-5-ylmethanol, are monoclinic, space group P2<sub>1</sub>/c, with a 15.5724(3), b 7.4976(2), c 24.2640(5) Å,  $\beta$  128.4980(10)°; Z = 4, dc = 1.381; R = 0.043, Rw(F<sub>2</sub>) = 0.116 for 3888 reflections. The imidazole and tetrazole rings are at angles of 85.0(2) and 51.8(1)°, resp., to the Ph rings to which they are attached, while the dihedral angle between the latter two rings is 46.7(1)°. The coordination sphere of the metal cation consists of six tetrazolyl N atoms, the MeOH O atom and the  $\pi$  cloud of one of the Ph rings. These interactions determine the formation of columns of mol. anions that lie parallel to the b axis, while H bonding contributes to intercolumnar cohesion. Far from the center of the columns, the hydrocarbon chain is immersed in a hydrophobic environment.

REFERENCE COUNT: 17 THERE ARE 17 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 22 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2002:207516 CAPLUS

DOCUMENT NUMBER: 136:236883

TITLE: Blood pressure-lowering compositions containing caffeic acid derivatives

INVENTOR(S): Suzuki, Atsushi; Ochiai, Ryuji; Kagawa, Taiji; Tokimitsu, Ichiro

PATENT ASSIGNEE(S): Kao Corp., Japan

SOURCE: Jpn. Kokai Tokkyo Koho, 5 pp.

CODEN: JKXXAF

DOCUMENT TYPE: Patent

LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 2002080354	A2	20020319	JP 2000-268099	20000905
PRIORITY APPLN. INFO.:			JP 2000-268099	20000905

AB This invention relates to blood pressure-lowering compns. containing (1) caffeic acid, chlorogenic acid, ferulic acid, and/or esters and salts thereof and (2) antihypertensives. For example, soft capsules were formulated containing caffeic acid 5, enalapril 5, corn starch 44, crystalline cellulose 40, CaCMC 5, silica 0.5, and Mg stearate 0.5 %.

L9 ANSWER 23 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2001:798216 CAPLUS

DOCUMENT NUMBER: 135:344489

TITLE: Detritylation process for the synthesis of losartan potassium using potassium hydroxide and a C1-4 alkanol solvent

INVENTOR(S): Fischer, Janos; Ballo, Ildiko; Petenyi, Endrene; Kreidl, Janos; Czibula, Laszlo; Nemes, Andras; Deutsch Juhasz, Ida; Werk Papp, Eva; Nagy Bagdy, Judit; Hegedus, Istvan; Farkas, Jenome

PATENT ASSIGNEE(S): Richter Gedeon Vegyeszeti Gyar Rt., Hung.

SOURCE: PCT Int. Appl., 15 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001081336	A1	20011101	WO 2001-HU47	20010420
WO 2001081336	C1	20020829		
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
AU 2001054998	A5	20011107	AU 2001-54998	20010420
EP 1274702	A1	20030115	EP 2001-928134	20010420
EP 1274702	B1	20040211		
R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR			
JP 2003531203	T2	20031021	JP 2001-578426	20010420
EE 200200460	A	20031215	EE 2002-460	20010420
AT 259366	E	20040215	AT 2001-928134	20010420
ES 2215130	T3	20041001	ES 2001-1928134	20010420
US 2003078435	A1	20030424	US 2002-182109	20020724
US 6710183	B2	20040323		
BG 107031	A	20030829	BG 2002-107031	20020823
PRIORITY APPLN. INFO.:			HU 2000-1618	A 20000421
			WO 2001-HU47	W 20010420

OTHER SOURCE(S): CASREACT 135:344489; MARPAT 135:344489

AB Losartan potassium (m.p. 262-264°) is prepared in high yield and selectivity by reacting the corresponding tritylated derivative [e.g., 2-butyl-4-chloro-1-[[2'-(2-triphenylmethyl-2H-tetrazol-5-yl)-1,1'-biphenyl-4-yl]methyl]-1H-imidazole-4-methanol] in an C1-4 alkanol (e.g., methanol) solvent with 0.1-1 equiv of potassium hydroxide and isolating the product after crystallizing out by changing the solvent to an aprotic (e.g., acetonitrile) or weakly protic solvent.

REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS

## RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 24 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2001:338762 CAPLUS

DOCUMENT NUMBER: 134:362292

TITLE: Methods of determining individual hypersensitivity to a pharmaceutical agent from gene expression profile

INVENTOR(S): Farr, Spencer

PATENT ASSIGNEE(S): Phase-1 Molecular Toxicology, USA

SOURCE: PCT Int. Appl., 222 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001032928	A2	20010510	WO 2000-US30474	20001103
WO 2001032928	A3	20020725		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				

PRIORITY APPLN. INFO.: US 1999-165398P P 19991105  
US 2000-196571P P 20000411

AB The invention discloses methods, gene databases, gene arrays, protein arrays, and devices that may be used to determine the hypersensitivity of individuals to a given agent, such as drug or other chemical, in order to prevent toxic side effects. In one embodiment, methods of identifying hypersensitivity in a subject by obtaining a gene expression profile of multiple genes associated with hypersensitivity of the subject suspected to be hypersensitive, and identifying in the gene expression profile of the subject a pattern of gene expression of the genes associated with hypersensitivity are disclosed. The gene expression profile of the subject may be compared with the gene expression profile of a normal individual and a hypersensitive individual. The gene expression profile of the subject that is obtained may comprise a profile of levels of mRNA or cDNA. The gene expression profile may be obtained by using an array of nucleic acid probes for the plurality of genes associated with hypersensitivity. The expression of the genes predetd. to be associated with hypersensitivity is directly related to prevention or repair of toxic damage at the tissue, organ or system level. Gene databases arrays and apparatus useful for identifying hypersensitivity in a subject are also disclosed.

L9 ANSWER 25 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1999:45216 CAPLUS

DOCUMENT NUMBER: 130:115010

TITLE: Process for the crystallization of losartan

INVENTOR(S): Breen, Patrick; Dienemann, Erik A.; Epstein, Albert D.; Larson, Karen A.; Kennedy, Michael T.; Mahadevan, Hari

PATENT ASSIGNEE(S): Merck and Co., Inc., USA

SOURCE: U.S., 7 pp.  
 CODEN: USXXAM  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 5859258	A	19990112	US 1997-959209	19971028
PRIORITY APPLN. INFO.:			US 1997-959209	19971028

AB Losartan potassium (I) is an angiotensin II antagonist useful in the treatment of hypertension and congestive heart failure. This invention relates to the process for the controlled crystallization of losartan potassium utilizing anti-solvent addition combined with massive seeding in order to obtain the desired crystal morphol. and bulk phys. properties necessary for successful formulation. Isopropanol 25.4 kg and 8.0 kg I were charged to a vessel along with 930 mL of distilled water. In a sep. vessel, 12.4 kg cyclohexane and 40 g I milled seed were heated to 60-65° and added to the above vessel until the solution became cloudy. The KF (Karl Fischer titration) at which the cloud point occurred was 1.90 % and the amount of cyclohexane slurry used to reach the cloud point was 6.2 kg. The batch was then seeded with 400 g finely-milled I and aged at reflux (70°) for 1 h. The batch was distilled at constant volume with simultaneous addition of 35 kg of 75:25 cyclohexane:isopropanol to achieve a batch KF of 0.54%. Distillates were collected with addition of 6 kg of cyclohexane to the batch during the concentration step. The batch was filtered under a N atmospheric and the cake was washed with 20 kg of 75:25 cyclohexane:isopropanol followed by 20 kg of cyclohexane. The batch was dried on trays at 45-50° under vacuum to obtain highly purified crystals of I.

REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 26 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN  
 ACCESSION NUMBER: 1998:293497 CAPLUS  
 DOCUMENT NUMBER: 128:326548  
 TITLE: Process for the crystallization of losartan  
 INVENTOR(S): Breen, Patrick; Dienemann, Erik A.; Epstein, Albert D.; Larson, Karen A.; Kennedy, Michael T.; Mahadevan, Hari  
 PATENT ASSIGNEE(S): Merck & Co., Inc., USA; Breen, Patrick; Dienemann, Erik A.; Epstein, Albert D.; Larson, Karen A.; Kennedy, Michael T.; Mahadevan, Hari  
 SOURCE: PCT Int. Appl., 20 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9818787	A1	19980507	WO 1997-US19442	19971024
W:	AL, AM, AU, AZ, BA, BB, BG, BR, BY, CA, CN, CU, EE, GE, HU, ID, IL, IS, JP, KG, KR, KZ, LC, LK, LR, LT, LV, MD, MG, MK, MN, MX, NO, NZ, PL, RO, RU, SG, SI, SK, SL, TJ, TM, TR, TT, UA, US,			

UZ, VN, YU  
 RW: GH, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR,  
 GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA,  
 GN, ML, MR, NE, SN, TD, TG

AU 9850898	A1	19980522	AU 1998-50898	19971024
EP 937068	A1	19990825	EP 1997-913800	19971024
EP 937068	B1	20020313		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, PT, IE, FI, RO				
BR 9712390	A	19990831	BR 1997-12390	19971024
CN 1241186	A	20000112	CN 1997-180909	19971024
CN 1101393	B	20030212		
JP 2000504343	T2	20000411	JP 1998-520675	19971024
JP 3249827	B2	20020121		
AT 214388	E	20020315	AT 1997-913800	19971024
PT 937068	T	20020731	PT 1997-913800	19971024
ES 2173433	T3	20021016	ES 1997-913800	19971024
SK 282875	B6	20030109	SK 1999-570	19971024
HR 970565	B1	20030228	HR 1997-970565	19971024
CZ 291672	B6	20030416	CZ 1999-1515	19971024
TW 411338	B	20001111	TW 1997-86116083	19971029

## PRIORITY APPLN. INFO.:

US 1996-29326P	P	19961029
GB 1996-25804	A	19961212
US 1996-29326	P	19961029
WO 1997-US19442	W	19971024

AB Losartan potassium is an angiotensin II antagonist  
 useful in the treatment of hypertension and congestive heart failure.  
 This invention relates to the process for the controlled crystallization  
 of losartan potassium utilizing anti-solvent addition  
 combined with massive seeding in order to obtain the desired  
 crystal morphol. and bulk phys. properties necessary for  
 successful formulation.

REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS  
 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 27 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1995:890142 CAPLUS

DOCUMENT NUMBER: 123:313978

TITLE: Polymorphs of losartan  
 potassium and a process for the preparation of  
 polymorph forms I and II of losartan  
 potassium

INVENTOR(S): Campbell, Gordon Creston, Jr.; Dwivedi, Anil M.;  
 Levorse, Dorothy A.; McCauley, James A.; Raghavan,  
 Krishnaswamy S.

PATENT ASSIGNEE(S): Merck and Co., Inc., USA; du Pont de Nemours, E. I.,  
 and Co.; Dupont Merck Pharmaceutical Co.

SOURCE: PCT Int. Appl., 54 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

## PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9517396	A1	19950629	WO 1994-US14768	19941221
W: AM, AU, BB, BG, BR, BY, CA, CN, CZ, EE, FI, GE, HU, JP, KG, KR, KZ, LK, LR, LT, LV, MD, MG, MN, NO, NZ, PL, RO, RU, SI, SK, TJ, TT, UA, US, UZ				

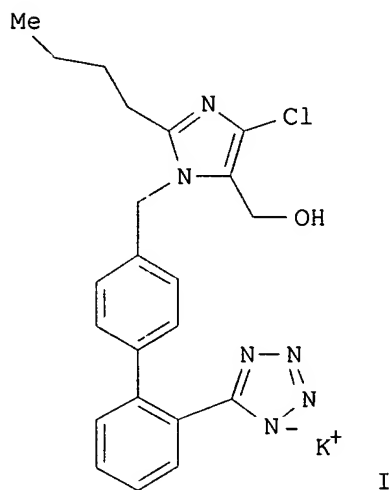


Andrew Freistein 10/524,993

RW: KE, MW, SD, SZ, AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU,  
MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN,  
TD, TG

CA 2179067	AA	19950629	CA 1994-2179067	19941221
AU 9514058	A1	19950710	AU 1995-14058	19941221
AU 685898	B2	19980129		
EP 736021	A1	19961009	EP 1995-905449	19941221
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, NL, PT, SE				
JP 09507075	T2	19970715	JP 1994-517594	19941221
US 5608075	A	19970304	US 1995-371937	19950112
PRIORITY APPLN. INFO.:			US 1993-173440	A 19931223
			WO 1994-US14768	W 19941221

GI



AB Polymorphic forms of losartan potassium, I, a known angiotensin II-inhibiting antihypertensive, are prepared Numerous spectral, thermal, and X-ray data of I form I and II are reported, and I-containing formulations are presented along with angiotensin II receptor inhibition data.

=>

---Logging off of STN---

=>

Executing the logoff script...

=> LOG Y

COST IN U.S. DOLLARS	SINCE FILE	TOTAL
	ENTRY	SESSION
FULL ESTIMATED COST	88.44	255.59
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	SINCE FILE	TOTAL
	ENTRY	SESSION

Andrew Freistein 10/524,993

CA SUBSCRIBER PRICE

-20.25

-20.25

STN INTERNATIONAL LOGOFF AT 11:23:49 ON 16 AUG 2006